

```

chain nodes :
  10  11  12  20  21  22  23  24
ring nodes :
  1  2  3  4  5  6  7  8  9  13  14  15  16  17  18
chain bonds :
  1-20  2-11  4-10  11-12  12-17  20-21  21-22  21-23  23-24
ring bonds :
  1-2  1-6  2-3  3-4  4-5  5-6  5-7  6-9  7-8  8-9  13-14  13-18  14-15
  15-16  16-17  17-18
exact/norm bonds :
  1-2  1-6  1-20  2-3  2-11  3-4  4-5  4-10  5-6  5-7  6-9  7-8  8-9  11-12
  21-22  21-23  23-24
exact bonds :
  12-17  20-21
normalized bonds :
  13-14  13-18  14-15  15-16  16-17  17-18
isolated ring systems :
  containing 1 :

```

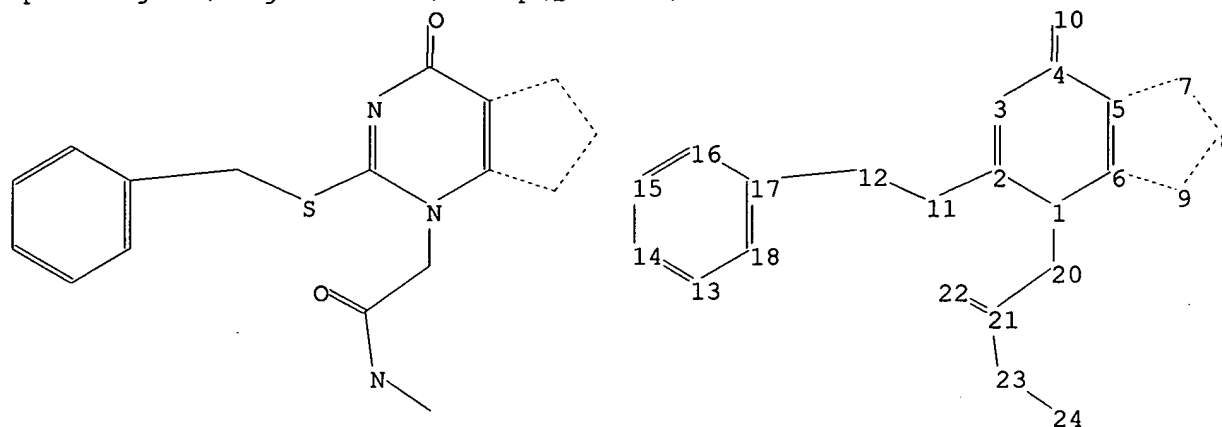
```

Match level :
  1:Atom  2:Atom  3:Atom  4:Atom  5:Atom  6:Atom  7:Atom  8:Atom  9:Atom
 10:CLASS 11:CLASS 12:CLASS 13:Atom 14:Atom 15:Atom 16:Atom 17:Atom
 18:Atom 20:CLASS 21:CLASS 22:CLASS 23:CLASS 24:CLASS

```

=>

Uploading C:\Program Files\Stnexp\Queries\10694561.str



chain nodes :

10 11 12 20 21 22 23 24

ring nodes :

1 2 3 4 5 6 7 8 9 13 14 15 16 17 18

chain bonds :

1-20 2-11 4-10 11-12 12-17 20-21 21-22 21-23 23-24

ring bonds :

1-2 1-6 2-3 3-4 4-5 5-6 5-7 6-9 7-8 8-9 13-14 13-18 14-15 15-16 16-17 17-18

exact/norm bonds :

1-2 1-6 1-20 2-3 2-11 3-4 4-5 4-10 5-6 5-7 6-9 7-8 8-9 11-12 21-22 21-23 23-24

exact bonds :

12-17 20-21

normalized bonds :

13-14 13-18 14-15 15-16 16-17 17-18

isolated ring systems :

containing 1 :

Match level :

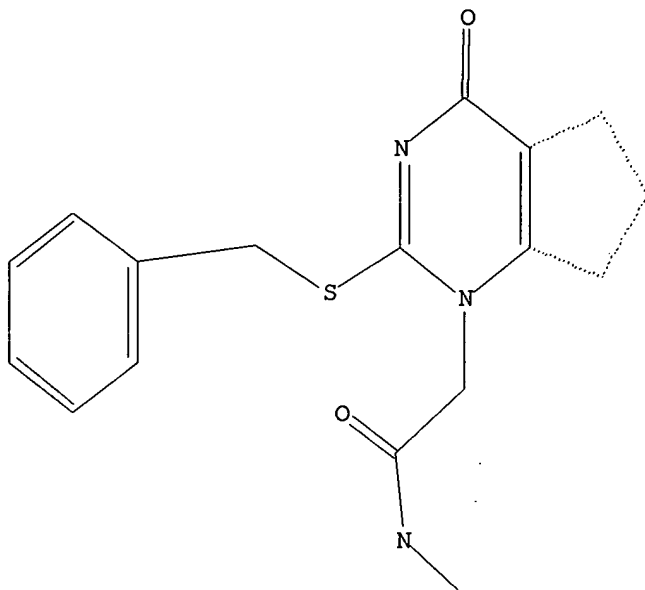
1:Atom 2:Atom 3:Atom 4:Atom 5:Atom 6:Atom 7:Atom 8:Atom 9:Atom 10:CLASS
 11:CLASS 12:CLASS 13:Atom 14:Atom 15:Atom 16:Atom 17:Atom 18:Atom 20:CLASS
 21:CLASS 22:CLASS 23:CLASS 24:CLASS

L1 STRUCTURE UPLOADED

=> d l1

L1 HAS NO ANSWERS

L1 STR



Structure attributes must be viewed using STN Express query preparation.

=> s l1 sss sam

SAMPLE SEARCH INITIATED 20:11:09 FILE 'REGISTRY'

SAMPLE SCREEN SEARCH COMPLETED - 34 TO ITERATE

100.0% PROCESSED 34 ITERATIONS

1 ANSWERS

SEARCH TIME: 00.00.01

FULL FILE PROJECTIONS: ONLINE **COMPLETE**

BATCH **COMPLETE**

PROJECTED ITERATIONS: 331 TO 1029

PROJECTED ANSWERS: 1 TO 80

L2 1 SEA SSS SAM L1

=> s l1 sss ful

FULL SEARCH INITIATED 20:11:17 FILE 'REGISTRY'

FULL SCREEN SEARCH COMPLETED - 612 TO ITERATE

100.0% PROCESSED 612 ITERATIONS

29 ANSWERS

SEARCH TIME: 00.00.01

L3 29 SEA SSS FUL L1

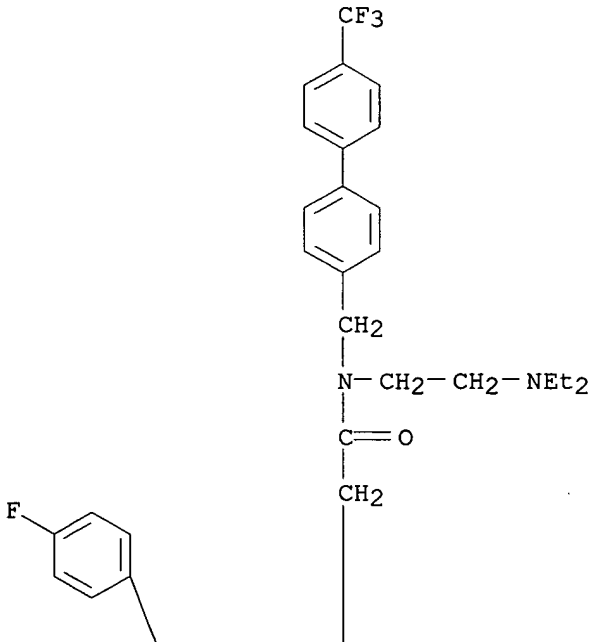
=> => s l3

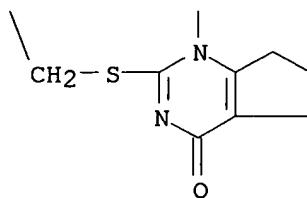
L4 6 L3

=> d l4 1-6 bib,ab,hitstr

L4 ANSWER 1 OF 6 CAPLUS COPYRIGHT 2004 ACS on STN
 AN 2004:720873 CAPLUS
 DN 141:342698
 TI SB-480848 (GlaxoSmithKline)
 AU Rotella, David P.
 CS Lexicon Pharmaceuticals, Princeton, NJ, 08540, USA
 SO Current Opinion in Investigational Drugs (Thomson Scientific) (2004),
 5(3), 348-351
 CODEN: COIDAZ; ISSN: 1472-4472
 PB Thomson Scientific
 DT Journal; General Review
 LA English
 AB A review. SB-480848 (synonyms/analogs: SB-435445, Lp-PLA2 inhibitor) is a reversible lipoprotein-associated phospholipase A2 inhibitor under development by GlaxoSmithKline for the potential treatment of atherosclerosis. Phase II trials with SB-480848 are currently underway.
 IT **356057-34-6P**, SB 480848
 RL: ADV (Adverse effect, including toxicity); DMA (Drug mechanism of action); PAC (Pharmacological activity); PKT (Pharmacokinetics); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)
 (SB 435445; reversible lipoprotein-associated phospholipase A2 inhibitor SB-480848 for potential treatment of atherosclerosis)
 RN 356057-34-6 CAPLUS
 CN 1H-Cyclopentapyrimidine-1-acetamide, N-[2-(diethylamino)ethyl]-2-[[4-(4-fluorophenyl)methyl]thio]-4,5,6,7-tetrahydro-4-oxo-N-[[4'-(trifluoromethyl)[1,1'-biphenyl]-4-yl]methyl]- (9CI) (CA INDEX NAME)

PAGE 1-A





RE.CNT 28 THERE ARE 28 CITED REFERENCES AVAILABLE FOR THIS RECORD
ALL CITATIONS AVAILABLE IN THE RE FORMAT

L4 ANSWER 2 OF 6 CAPLUS COPYRIGHT 2004 ACS on STN
 AN 2003:837075 CAPLUS
 DN 139:337982
 TI Preparation of pyridone and pyrimidone compounds as inhibitors of the enzyme Lp-PLA2
 IN Leach, Colin Andrew; Smith, Stephen Allan
 PA Glaxo Group Limited, UK
 SO PCT Int. Appl., 61 pp.
 CODEN: PIXXD2
 DT Patent
 LA English
 FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	WO 2003087088	A2	20031023	WO 2003-GB1550	20030410
	WO 2003087088	A3	20040108		
	W:	AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH, PL, PT, RO, RU, SD, SE, SG, SK, SL, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW			
	RW:	GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG			
PRAI	GB 2002-8280	A	20020410		

OS MARPAT 139:337982

AB The title compds. [I; R1 = (un)substituted aryl; R2 = halo, alkyl, alkoxy, etc.; R3 = H, halo, alkyl, hydroxyalkyl; R2 and R3 together with the pyridone or pyrimidine ring carbons to which they are attached form (un)substituted fused 5-6 membered carbocyclic ring, fused benzo or heteroaryl ring; R4 = (CH2)_n substituted by benzimidazole or 5-6 membered heteroaryl; R5 = (un)substituted (hetero)aryl; R6 = (un)substituted (hetero)aryl; X = CH, N; Y = alkylene, CH:CH, (CH2)_mS; n = 1-4; m = 1-2] that are inhibitors of the enzyme Lp-PLA2 and are of use in therapy, in particular for treating atherosclerosis, were prepared Thus, amidation of 2-[2-(2,3-difluorobenzylthio)-4-oxo-4H-quinolin-1-yl]acetic acid with N-[2-(1-methylimidazol-4-yl)ethyl]-4'-trifluoromethylbiphen-4-ylmethylamine (prepns. given) afforded the quinolinone II. The exemplified compds. I showed IC50 values in the range <0.1 to 100 nM against Lp-PLA2.

IT **615578-19-3P**

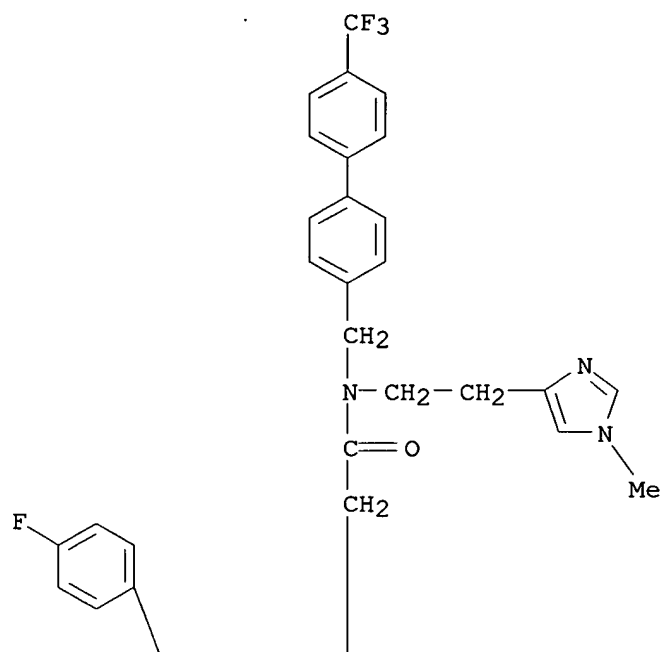
RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(preparation of pyridone compds. as inhibitors of the enzyme Lp-PLA2)

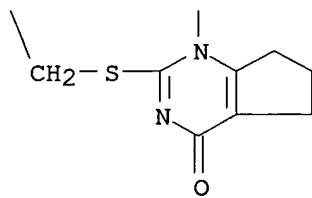
RN 615578-19-3 CAPLUS

CN 1H-Cyclopentapyrimidine-1-acetamide, 2-[[[4-fluorophenyl)methyl]thio]-4,5,6,7-tetrahydro-N-[2-(1-methyl-1H-imidazol-4-yl)ethyl]-4-oxo-N-[[4'-(trifluoromethyl)[1,1'-biphenyl]-4-yl]methyl]- (9CI) (CA INDEX NAME)

PAGE 1-A



PAGE 2-A



L4 ANSWER 3 OF 6 CAPLUS COPYRIGHT 2004 ACS on STN
 AN 2003:836853 CAPLUS
 DN 139:337978
 TI Preparation of N-substituted pyridinone and pyrimidinone derivatives for
 use as Lp-PLA2 inhibitors in the treatment of atherosclerosis
 IN Leach, Colin Andrew; Smith, Stephen Allan
 PA Glaxo Group Limited, UK
 SO PCT Int. Appl., 38 pp.
 CODEN: PIXXD2
 DT Patent
 LA English
 FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	WO 2003086400	A1	20031023	WO 2003-GB1544	20030410
	W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH, PL, PT, RO, RU, SD, SE, SG, SK, SL, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG				
PRAI	GB 2002-8279	A	20020410		

OS MARPAT 139:337978

AB The title compds. [I; R1 = (un)substituted aryl; R2 = halo, alkyl, alkoxy, etc.; R3 = H, halo, alkyl, hydroxyalkyl; R2 and R3 together with the pyridone or pyrimidinone ring carbons to which they are attached form (un)substituted fused 5-6 membered carbocyclic ring, fused benzo or heteroaryl ring; R4 = alkyl substituted by 5-7 membered saturated heterocyclyl comprising N and optionally O or S; R5 = (un)substituted (hetero)aryl; R6 = (un)substituted (hetero)aryl; X = CH, N; Y = alkylene, CH:CH, (CH₂)_nS; n = 1-3] that are inhibitors of the enzyme Lp-PLA2 and are of use in therapy, in particular for treating atherosclerosis, were prepared. Thus, amidation of 2-[2-(2,3-difluorobenzylthio)-4-oxo-4H-quinolin-1-yl]acetic acid with N-(1-thiazol-2-ylmethylpiperidin-4-yl)-4'-trifluoromethylbiphen-4-ylmethylamine (preps. given) afforded the quinolinone II. The exemplified compds. I showed IC₅₀ values in the range <0.1 to 100 nM against Lp-PLA2.

IT **615577-22-5P**

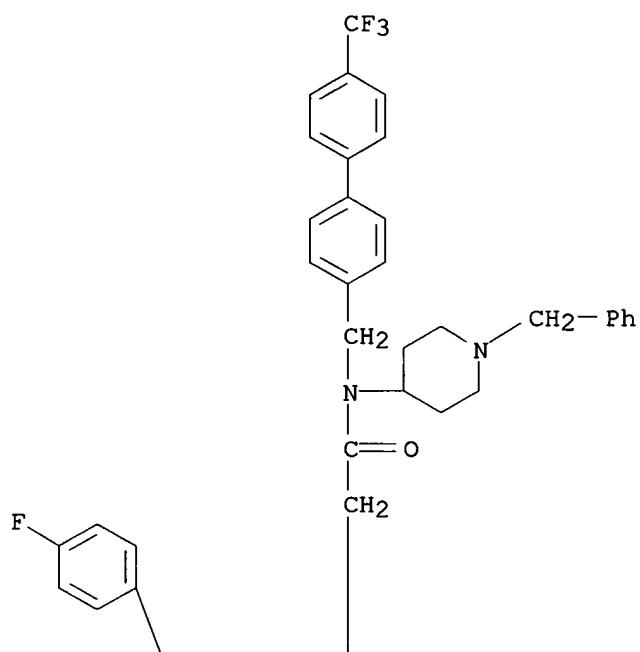
RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(preparation of pyridinone and pyrimidinone derivs. for use as Lp-PLA2 inhibitors in the treatment of atherosclerosis)

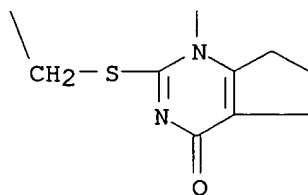
RN 615577-22-5 CAPLUS

CN 1H-Cyclopentapyrimidine-1-acetamide, 2-[[[(4-fluorophenyl)methyl]thio]-4,5,6,7-tetrahydro-4-oxo-N-[1-(phenylmethyl)-4-piperidinyl]-N-[[4'-(trifluoromethyl)[1,1'-biphenyl]-4-yl]methyl]- (9CI) (CA INDEX NAME)

PAGE 1-A



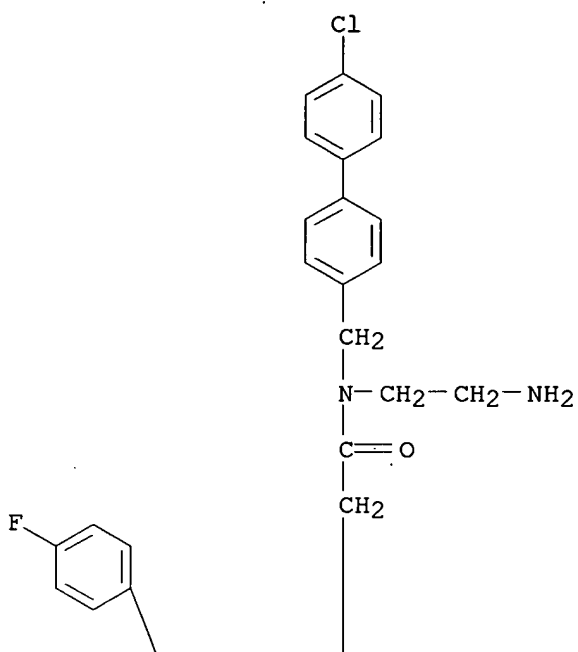
PAGE 2-A



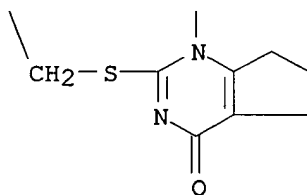
RE.CNT 4 THERE ARE 4 CITED REFERENCES AVAILABLE FOR THIS RECORD
ALL CITATIONS AVAILABLE IN THE RE FORMAT

L4 ANSWER 4 OF 6 CAPLUS COPYRIGHT 2004 ACS on STN
AN 2003:215748 CAPLUS
DN 139:78433
TI The identification of clinical candidate SB-480848: a potent inhibitor of
lipoprotein-associated phospholipase A2
AU Blackie, Josie A.; Bloomer, Jackie C.; Brown, Murray J. B.; Cheng,
Hung-Yuan; Hammond, Beverley; Hickey, Deirdre M. B.; Ife, Robert J.;
Leach, Colin A.; Lewis, V. Ann; Macphee, Colin H.; Milliner, Kevin J.;
Moores, Kitty E.; Pinto, Ivan L.; Smith, Stephen A.; Stansfield, Ian G.;
Stanway, Steven J.; Taylor, Maxine A.; Theobald, Colin J.
CS Medicines Research Centre, GlaxoSmithKline, Stevenage, SG1 2NY, UK
SO Bioorganic & Medicinal Chemistry Letters (2003), 13(6), 1067-1070
CODEN: BMCLE8; ISSN: 0960-894X
PB Elsevier Science B.V.
DT Journal
LA English
OS CASREACT 139:78433
AB Modification of the pyrimidone 5-substituent in clin. candidate SB-435495
has given a series of inhibitors of recombinant lipoprotein-associated
phospholipase A2 with sub-nanomolar potency. Cyclopentyl fused derivative 21,
SB-480848, showed an enhanced in vitro and in vivo profile vs. SB-435495
and has been selected for progression to man.
IT **552857-62-2P 552857-63-3P**
RL: PAC (Pharmacological activity); PKT (Pharmacokinetics); SPN (Synthetic
preparation); THU (Therapeutic use); BIOL (Biological study); PREP
(Preparation); USES (Uses)
(design and structure activity of lipoprotein-associated phospholipase A2
inhibitor SB-480848)
RN 552857-62-2 CAPLUS
CN 1H-Cyclopentapyrimidine-1-acetamide, N-(2-aminoethyl)-N-[(4'-chloro[1,1'-
biphenyl]-4-yl)methyl]-2-[[(4-fluorophenyl)methyl]thio]-4,5,6,7-tetrahydro-
4-oxo- (9CI) (CA INDEX NAME)

PAGE 1-A

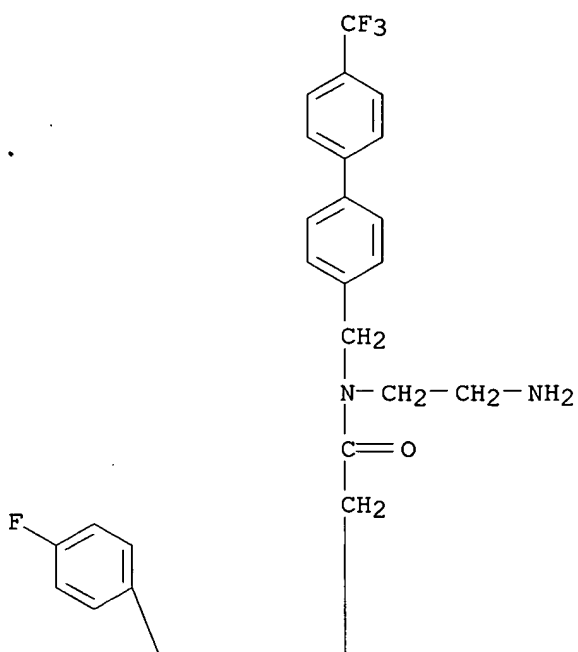


PAGE 2-A

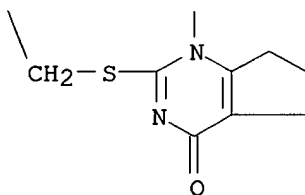


RN 552857-63-3 CAPLUS
 CN 1H-Cyclopentapyrimidine-1-acetamide, N-(2-aminoethyl)-2-[[(4-fluorophenyl)methyl]thio]-4,5,6,7-tetrahydro-4-oxo-N-[[4'-(trifluoromethyl)[1,1'-biphenyl]-4-yl]methyl]- (9CI) (CA INDEX NAME)

PAGE 1-A



PAGE 2-A



RE.CNT 9 THERE ARE 9 CITED REFERENCES AVAILABLE FOR THIS RECORD
ALL CITATIONS AVAILABLE IN THE RE FORMAT

L4 ANSWER 5 OF 6 CAPLUS COPYRIGHT 2004 ACS on STN
 AN 2003:154411 CAPLUS
 DN 138:187787

TI Novel processes for the preparation of pyrimidinone derivatives, useful as
 Lp-PLA2 inhibitors, and intermediates thereof

IN Mulholland, Keith Raymond; Ross, Andrew R.; Slater, Graham Ralph; Smith,
 Gillian Elizabeth

PA Smithkline Beecham PLC, UK

SO PCT Int. Appl., 14 pp.

CODEN: PIXXD2

DT Patent

LA English

FAN.CNT 1

Common Assignee

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	WO 2003016287	A2	20030227	WO 2002-EP9067	20020813
	WO 2003016287	A3	20031016		
	W:				
	AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN,				
	CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH,				
	GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR,				
	LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH,				
	PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TN, TR, TT, TZ,				
	UA, UG, US, UZ, VN, YU, ZA, ZM, ZW				
	RW:				
	GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY,				
	KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES,				
	FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, SK, TR, BF, BJ, CF,				
	CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG				
	EP 1456183	A2	20040915	EP 2002-794787	20020813
	R:				
	AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,				
	IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, SK				
	US 2004242875	A1	20041202	US 2004-485972	20040702
PRAI	GB 2001-19795	A	20010814		
	WO 2002-EP9067	W	20020813		

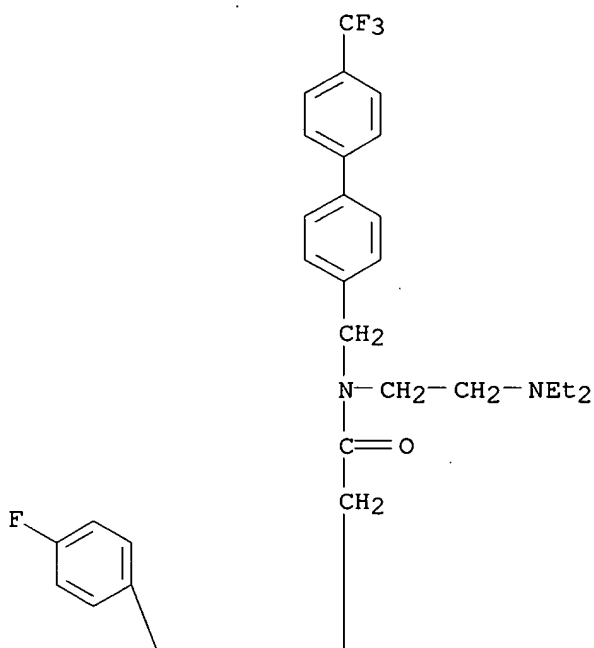
Check

OS CASREACT 138:187787; MARPAT 138:187787

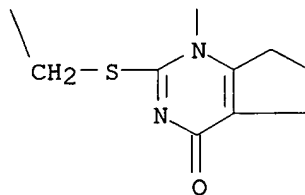
AB The invention relates to a process for the preparation of certain pyrimidinone compds., including the intermediates I and II [wherein: RaRb = (CH₂)₃₋₄; R1 = Ph optionally substituted by halogen], and the final target compds. III [wherein: RaRb = atoms to form 5-membered carbocyclic ring; R1 = 4-fluorophenyl; R2 = C1-3 alkyl substituted by NR₅R₆; R2 = Het-CO-2-alkyl; Het = 5- to 7-membered N-heterocyclyl with N optionally substituted by C1-6 alkyl; R3R4 = 4-[4-(trifluoromethyl)phenyl]phenyl; R5, R6 = H, C1-6 alkyl]. Compds. III, described in WO 01/60805, are known inhibitors (no data) of lipoprotein-associated phospholipase A2 (Lp-PLA2), useful, e.g., for prevention of acute coronary events caused by atherosclerosis. The literature methods of preparing III suffer from moderate yields due to poor selectivity in the alkylation of the pyrimidinone nucleus. The invention method gives selective N1-alkylation of the pyrimidinone nucleus, and does not require isolation of an intermediate ester, thus giving high yields and efficiency. For instance, cyclocondensation of Et 2-oxocyclopentanecarboxylate with thiourea in the presence of DBU gave 67.6% 5,6-trimethylene-2-thiouracil, which underwent S-alkylation by 4-fluorobenzyl chloride in the presence of K₂CO₃ and KI in Me₂CO to give 86.5% intermediate II [R1 = 4-FC₆H₄, RaRb = (CH₂)₃]. This compound was selectively O-silylated by (Me₃Si)₂NH and saccharin in CH₂Cl₂, selectively N1-alkylated by CF₃SO₂OCH₂CO₂Me, and then hydrolyzed directly by aqueous NaOH in iso-PrOH, to give 69% I [R1 = 4-FC₆H₄, RaRb = (CH₂)₃]. Amidation of this acid with the corresponding amine using DIPEA and TBTU in CH₂Cl₂, followed by recrystn. from iso-PrOAc, gave 88% target compound IV.

IT **356057-34-6P**, 1-[[[N-[2-(Diethylamino)ethyl]-N-[4-[4-(trifluoromethyl)phenyl]benzyl]amino]carbonyl]methyl]-2-(4-fluorobenzylthio)-5,6-trimethylenepyrimidin-4-one
 RL: IMF (Industrial manufacture); SPN (Synthetic preparation); PREP (Preparation)
 (product; processes for preparation of pyrimidinone derivs. useful as Lp-PLA2 inhibitors and their intermediates)
 RN 356057-34-6 CAPLUS
 CN 1H-Cyclopentapyrimidine-1-acetamide, N-[2-(diethylamino)ethyl]-2-[[4-(4-fluorophenyl)methyl]thio]-4,5,6,7-tetrahydro-4-oxo-N-[[4'-(trifluoromethyl)[1,1'-biphenyl]-4-yl]methyl]- (9CI) (CA INDEX NAME)

PAGE 1-A



PAGE 2-A



L4 ANSWER 6 OF 6 CAPLUS COPYRIGHT 2004 ACS on STN
 AN 2001:617985 CAPLUS
 DN 135:195570
 TI Preparation of pyrimidine-4-one derivatives as LDL-PLA2 inhibitors
 IN Hickey, Deirdre Mary Bernadette; Ife, Robert John; Leach, Colin Andrew;
 Pinto, Ivan Leo; Smith, Stephen Allan; Stanway, Steven James
 PA Smithkline Beecham P.L.C., UK
 SO PCT Int. Appl., 54 pp.
 CODEN: PIXXD2
 DT Patent
 LA English
 FAN.CNT 1

Appl PCS

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	WO 2001060805	A1	20010823	WO 2001-EP1515	20010213
	W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM				
	RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG				
	CA 2400554	AA	20010823	CA 2001-2400554	20010213
	AU 2001035466	A5	20010827	AU 2001-35466	20010213
	EP 1263740	A1	20021211	EP 2001-907522	20010213
	R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR				
	BR 2001008396	A	20030311	BR 2001-8396	20010213
	JP 2003523335	T2	20030805	JP 2001-560190	20010213
	NZ 520752	A	20040326	NZ 2001-520752	20010213
	RU 2235722	C2	20040910	RU 2002-124611	20010213
	US 2002103213	A1	20020801	US 2001-782930	20010214
	TW 550259	B	20030901	TW 2001-90103332	20010215
	NO 2002003828	A	20020930	NO 2002-3828	20020813
	ZA 2002006528	A	20030313	ZA 2002-6528	20020815
	BG 107034	A	20030430	BG 2002-107034	20020826
	US 6649619	B1	20031118	US 2003-357238	20030203
	US 2004097525	A1	20040520	US 2003-694561	20031027
PRAI	GB 2000-3636	A	20000216		
	GB 2001-1437	A	20010119		
	WO 2001-EP1515	W	20010213		
	US 2001-782930	B1	20010214		
	US 2003-357238	A3	20030203		

OS MARPAT 135:195570

AB The title compds. [I; Ra = H, halo, alkyl, etc.; Rb = H, halo, alkyl, etc.; Ra and Rb together = (CH₂)_n (n = 3-4) or Ra and Rb together with the pyrimidine ring carbon atoms to which they are attached form (un)substituted fused benzo or heteroaryl ring; Rc = H, alkyl; R2 = (un)substituted (hetero)aryl; R3 = H, alkyl, halo, etc.; R4 = (un)substituted (hetero)arylene; R5 = (un)substituted (hetero)aryl; n = 1-4; X = O, S; Y = (CH₂)_pOq (p = 1-3 and q = 0; p = 2-3 and q = 1); Z = O, a bond] which are inhibitors of the enzyme Lp-PLA2 useful in treating atherosclerosis, were prepared Thus, reacting N-[2-(diethylamino)ethyl]-4-(4-trifluoromethylphenyl)benzylamine with 1-(carboxymethyl)-2-(4-fluorobenzylthio)-5-ethylpyrimidin-4-one in the presence of HATU and (iso-Pr)₂NEt in CH₂Cl₂ afforded the pyrimidinone II. The compds. I

described in Examples were tested for Lp-PLA2 inhibition and showed IC50 values in the range <0.1 nM to 10 μ M.

IT **356057-38-0P**

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); RCT (Reactant); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); RACT (Reactant or reagent); USES (Uses)

(preparation of pyrimidine-4-one derivs. as LDL-PLA2 inhibitors)

RN 356057-38-0 CAPLUS

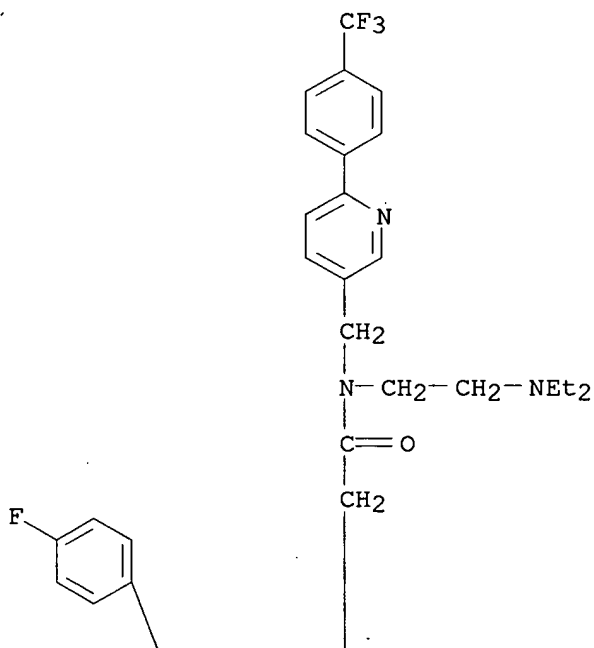
CN 1H-Cyclopentapyrimidine-1-acetamide, N-[2-(diethylamino)ethyl]-2-[[4-(4-fluorophenyl)methyl]thio]-4,5,6,7-tetrahydro-4-oxo-N-[[6-[4-(trifluoromethyl)phenyl]-3-pyridinyl]methyl]-, (2R,3R)-2,3-dihydroxybutanedioate (1:2) (9CI) (CA INDEX NAME)

CM 1

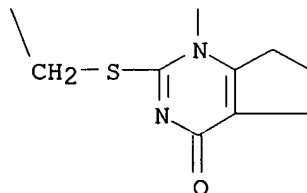
CRN 356057-37-9

CMF C35 H37 F4 N5 O2 S

PAGE 1-A



PAGE 2-A

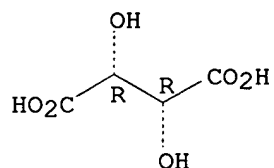


CM 2

CRN 87-69-4

CMF C4 H6 O6

Absolute stereochemistry.



IT 356057-34-6P 356057-35-7P 356057-36-8P
 356057-37-9P 356057-39-1P 356057-40-4P
 356057-69-7P 356057-87-9P 356057-88-0P
 356057-89-1P 356057-90-4P 356057-91-5P
 356057-92-6P 356057-93-7P 356057-94-8P
 356057-95-9P 356057-98-2P 356057-99-3P
 356058-00-9P 356058-03-2P 356058-05-4P
 356058-06-5P 356058-07-6P 356058-12-3P

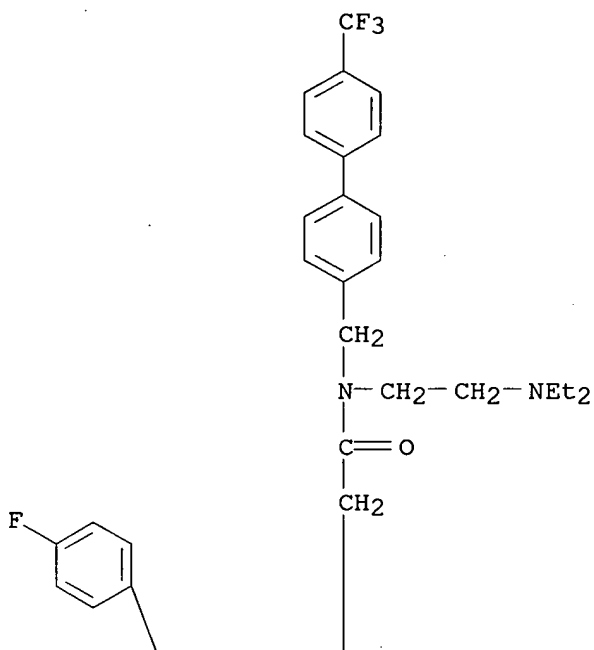
RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(preparation of pyrimidine-4-one derivs. as LDL-PLA2 inhibitors)

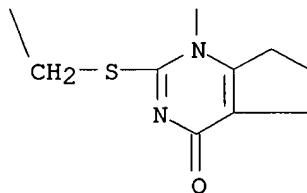
RN 356057-34-6 CAPLUS

CN 1H-Cyclopentapyrimidine-1-acetamide, N-[2-(diethylamino)ethyl]-2-[[4-fluorophenyl)methyl]thio]-4,5,6,7-tetrahydro-4-oxo-N-[[4'-(trifluoromethyl)[1,1'-biphenyl]-4-yl]methyl]- (9CI) (CA INDEX NAME)

PAGE 1-A



PAGE 2-A

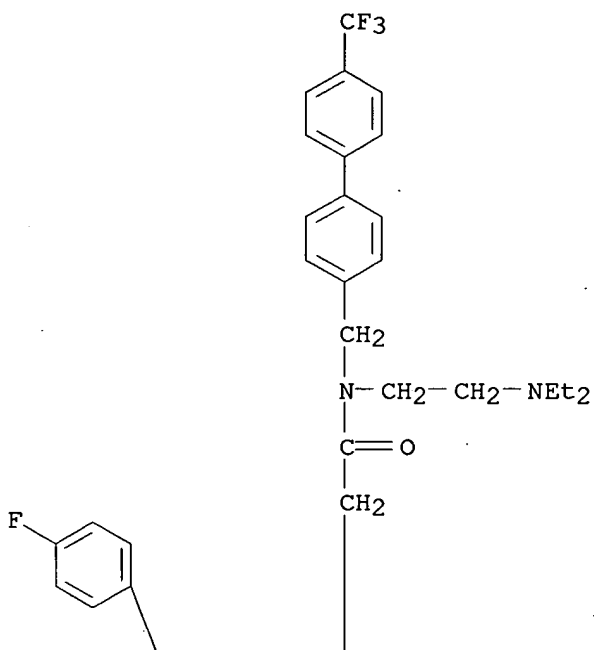


RN 356057-35-7 CAPLUS
 CN 1H-Cyclopentapyrimidine-1-acetamide, N-[2-(diethylamino)ethyl]-2-[[[4-fluorophenyl)methyl]thio]-4,5,6,7-tetrahydro-4-oxo-N-[[4'-(trifluoromethyl)[1,1'-biphenyl]-4-yl]methyl]-, (2R,3R)-2,3-dihydroxybutanedioate (1:2) (9CI) (CA INDEX NAME)

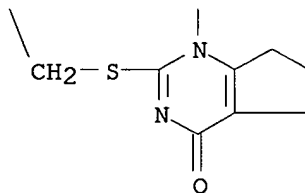
CM 1

CRN 356057-34-6
 CMF C36 H38 F4 N4 O2 S

PAGE 1-A



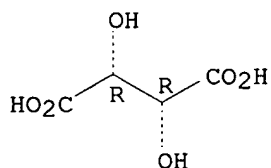
PAGE 2-A



CM 2

CRN 87-69-4
CMF C4 H6 O6

Absolute stereochemistry.



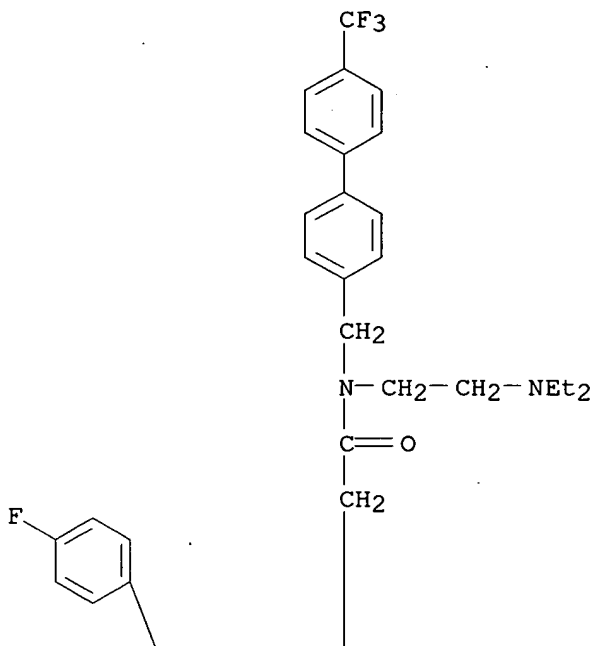
RN 356057-36-8 CAPLUS

CN 1H-Cyclopentapyrimidine-1-acetamide, N-[2-(diethylamino)ethyl]-2-[[4-

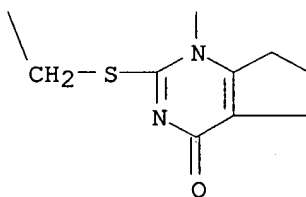
10/694,561

fluorophenyl)methyl]thio]-4,5,6,7-tetrahydro-4-oxo-N-[[4'-(trifluoromethyl)[1,1'-biphenyl]-4-yl]methyl]-, monohydrochloride (9CI)
(CA INDEX NAME)

PAGE 1-A



PAGE 2-A

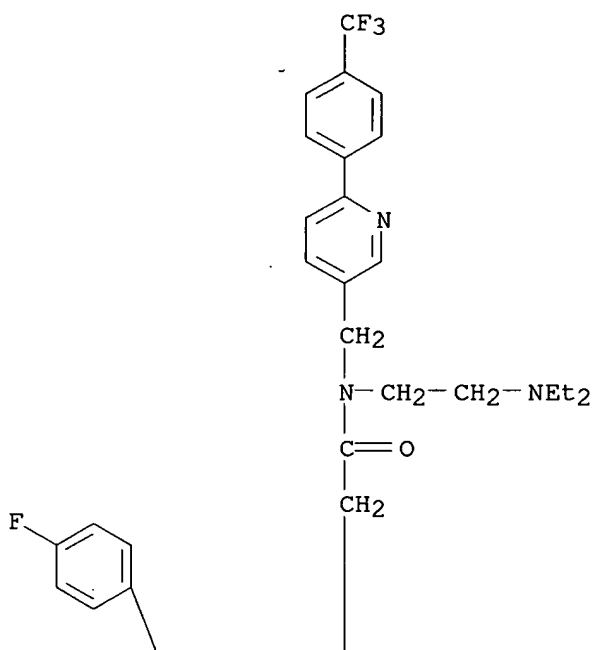


● HCl

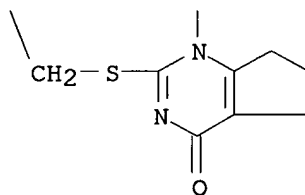
RN 356057-37-9 CAPLUS

CN 1H-Cyclopentapyrimidine-1-acetamide, N-[2-(diethylamino)ethyl]-2-[[4-(4-fluorophenyl)methyl]thio]-4,5,6,7-tetrahydro-4-oxo-N-[[6-[4-(trifluoromethyl)phenyl]-3-pyridinyl]methyl]- (9CI) (CA INDEX NAME)

PAGE 1-A

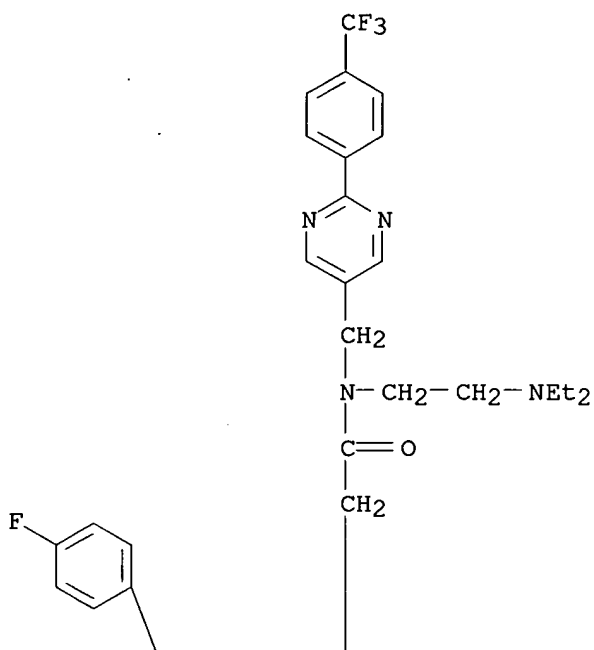


PAGE 2-A

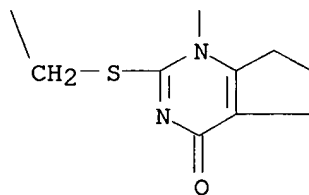


RN 356057-39-1 CAPLUS
 CN 1H-Cyclopentapyrimidine-1-acetamide, N-[2-(diethylamino)ethyl]-2-[[4-(4-fluorophenyl)methyl]thio]-4,5,6,7-tetrahydro-4-oxo-N-[[2-[4-(trifluoromethyl)phenyl]-5-pyrimidinyl]methyl]- (9CI) (CA INDEX NAME)

PAGE 1-A



PAGE 2-A

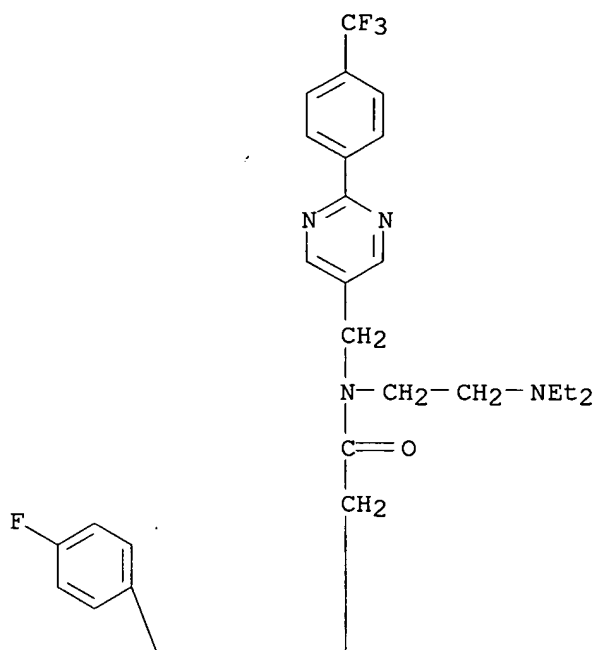


RN 356057-40-4 CAPLUS
 CN 1H-Cyclopentapyrimidine-1-acetamide, N-[2-(diethylamino)ethyl]-2-[[[4-(4-fluorophenyl)methyl]thio]-4,5,6,7-tetrahydro-4-oxo-N-[[2-[4-(trifluoromethyl)phenyl]-5-pyrimidinyl]methyl]-, (2R,3R)-2,3-dihydroxybutanedioate (1:2) (9CI) (CA INDEX NAME)

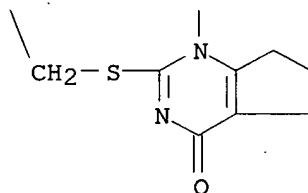
CM 1

CRN 356057-39-1
 CMF C34 H36 F4 N6 O2 S

PAGE 1-A



PAGE 2-A

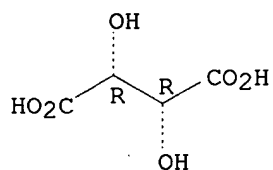


CM 2

CRN 87-69-4

CMF C4 H6 O6

Absolute stereochemistry.

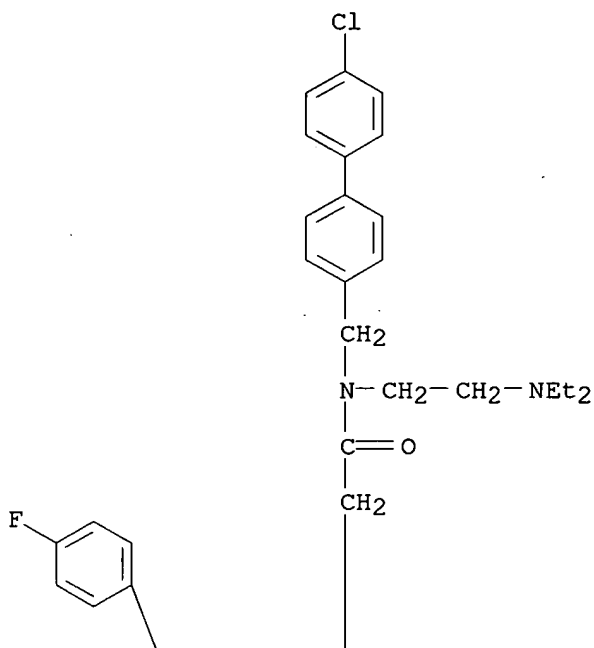


RN 356057-69-7 CAPLUS

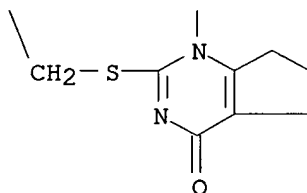
CN 1H-Cyclopentapyrimidine-1-acetamide, N-[(4'-chloro[1,1'-biphenyl]-4-

yl)methyl]-N-[2-(diethylamino)ethyl]-2-[[4-(4-fluorophenyl)methyl]thio]-
4,5,6,7-tetrahydro-4-oxo- (9CI) (CA INDEX NAME)

PAGE 1-A

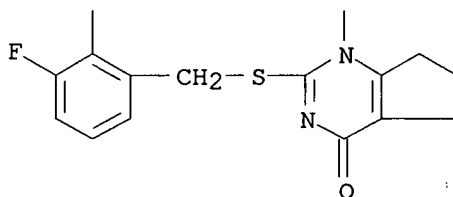
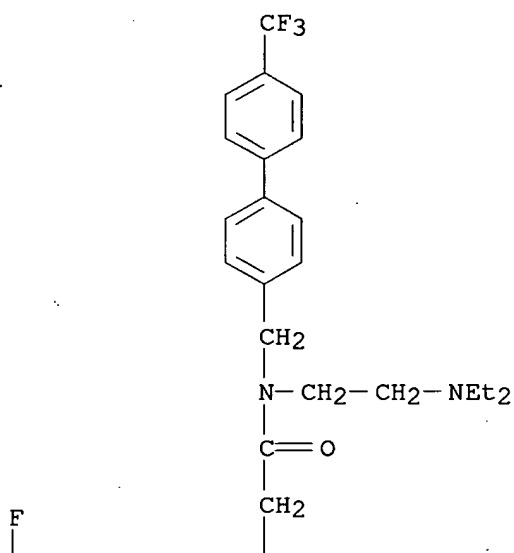


PAGE 2-A



RN 356057-87-9 CAPLUS

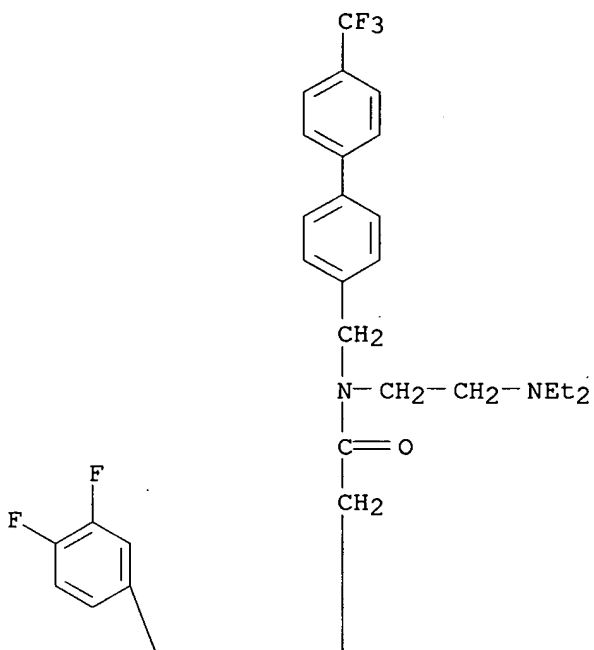
CN 1H-Cyclopentapyrimidine-1-acetamide, N-[2-(diethylamino)ethyl]-2-[[2,3-difluorophenyl)methyl]thio]-4,5,6,7-tetrahydro-4-oxo-N-[[4'-(trifluoromethyl)[1,1'-biphenyl]-4-yl]methyl]- (9CI) (CA INDEX NAME)



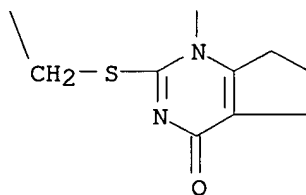
RN 356057-88-0 CAPLUS

CN 1H-Cyclopentapyrimidine-1-acetamide, N-[2-(diethylamino)ethyl]-2-[[(3,4-difluorophenyl)methyl]thio]-4,5,6,7-tetrahydro-4-oxo-N-[[4'-(trifluoromethyl)[1,1'-biphenyl]-4-yl]methyl]- (9CI) (CA INDEX NAME)

PAGE 1-A

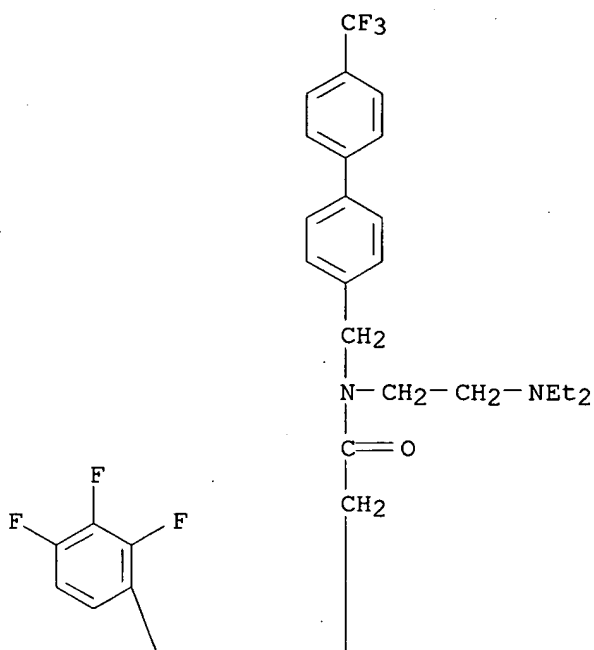


PAGE 2-A

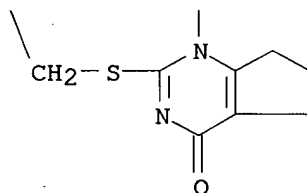


RN 356057-89-1 CAPLUS
 CN 1H-Cyclopentapyrimidine-1-acetamide, N-[2-(diethylamino)ethyl]-4,5,6,7-tetrahydro-4-oxo-N-[[4'-(trifluoromethyl)[1,1'-biphenyl]-4-yl]methyl]-2-[[2,3,4-trifluorophenyl]methyl]thio]- (9CI) (CA INDEX NAME)

PAGE 1-A

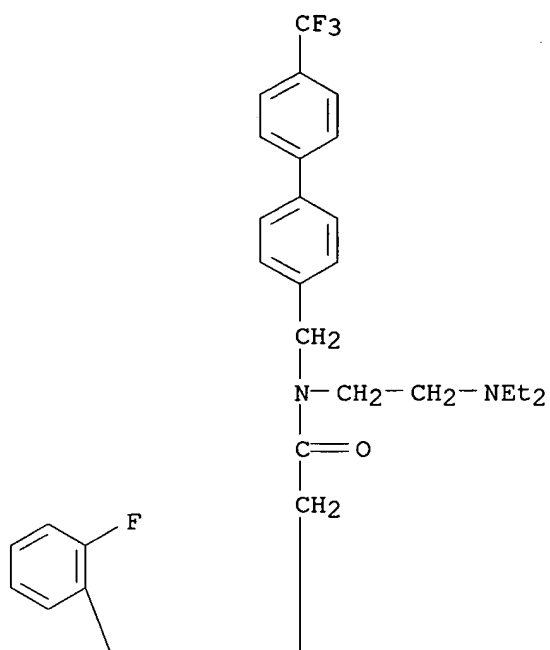


PAGE 2-A

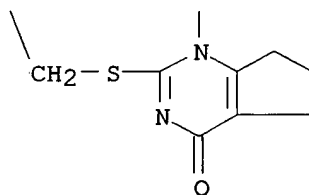


RN 356057-90-4 CAPLUS
 CN 1H-Cyclopentapyrimidine-1-acetamide, N-[2-(diethylamino)ethyl]-2-[[[2-fluorophenyl)methyl]thio]-4,5,6,7-tetrahydro-4-oxo-N-[[4'-(trifluoromethyl)[1,1'-biphenyl]-4-yl)methyl]- (9CI) (CA INDEX NAME)

PAGE 1-A



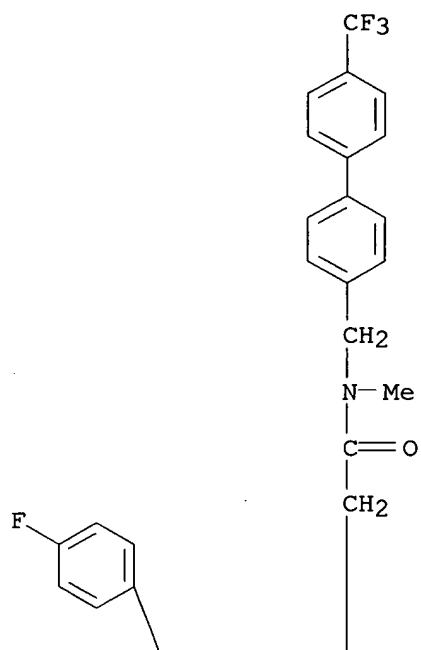
PAGE 2-A



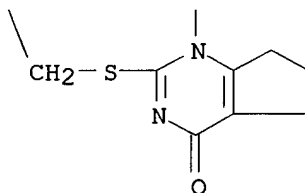
RN 356057-91-5 CAPLUS

CN 1H-Cyclopentapyrimidine-1-acetamide, 2-[[[4-fluorophenyl)methyl]thio]-
4,5,6,7-tetrahydro-N-methyl-4-oxo-N-[[4'-(trifluoromethyl)[1,1'-biphenyl]-
4-yl]methyl]- (9CI) (CA INDEX NAME)

PAGE 1-A

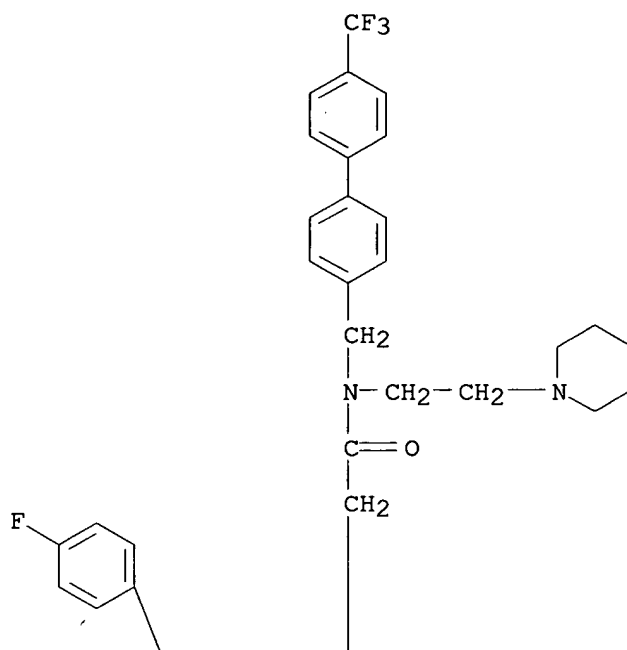


PAGE 2-A

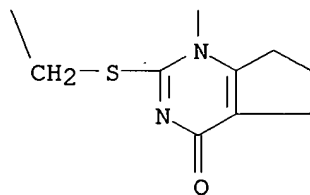


RN 356057-92-6 CAPLUS
 CN 1H-Cyclopentapyrimidine-1-acetamide, 2-[[[4-fluorophenyl)methyl]thio]-
 4,5,6,7-tetrahydro-4-oxo-N-[2-(1-piperidinyl)ethyl]-N-[[4'-
 (trifluoromethyl)[1,1'-biphenyl]-4-yl)methyl]- (9CI) (CA INDEX NAME)

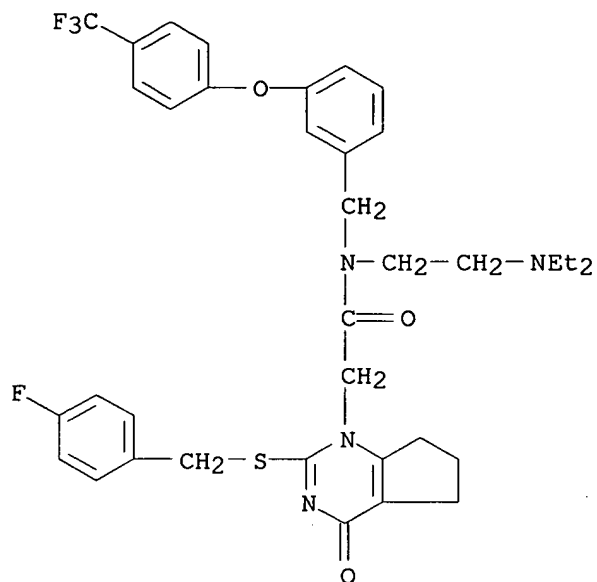
PAGE 1-A



PAGE 2-A



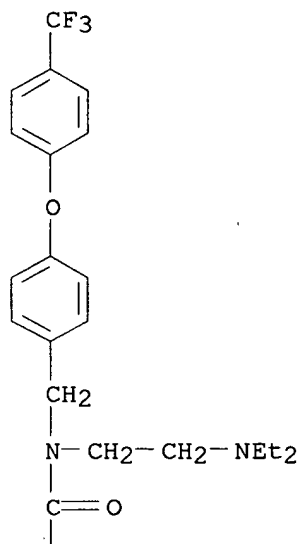
RN 356057-93-7 CAPLUS
 CN 1H-Cyclopentapyrimidine-1-acetamide, N-[2-(diethylamino)ethyl]-2-[[[4-fluorophenyl)methyl]thio]-4,5,6,7-tetrahydro-4-oxo-N-[[3-[4-(trifluoromethyl)phenoxy]phenyl)methyl]- (9CI) (CA INDEX NAME)



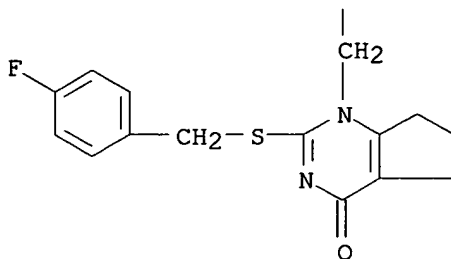
RN 356057-94-8 CAPLUS

CN 1H-Cyclopentapyrimidine-1-acetamide, N-[2-(diethylamino)ethyl]-2-[[4-(4-fluorophenyl)methyl]thio]-4,5,6,7-tetrahydro-4-oxo-N-[[4-[4-(trifluoromethyl)phenoxy]phenyl]methyl]- (9CI) (CA INDEX NAME)

PAGE 1-A

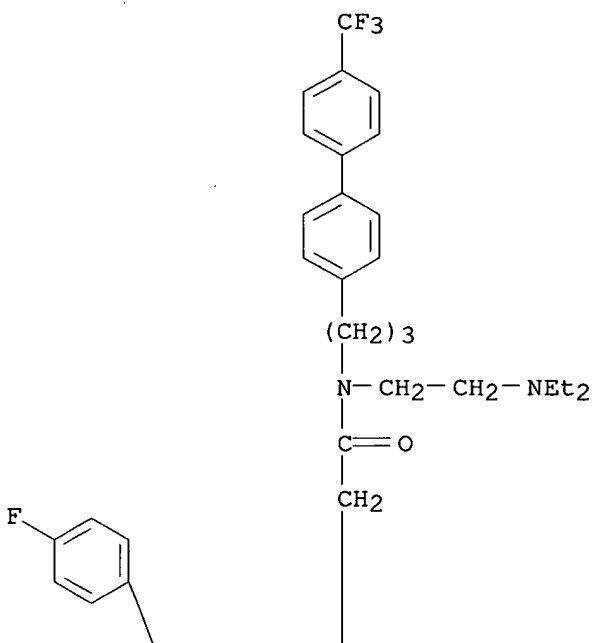


PAGE 2-A

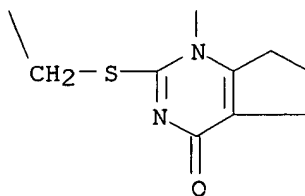


RN 356057-95-9 CAPLUS
 CN 1H-Cyclopentapyrimidine-1-acetamide, N-[2-(diethylamino)ethyl]-2-[[(4-fluorophenyl)methyl]thio]-4,5,6,7-tetrahydro-4-oxo-N-[3-[4'-(trifluoromethyl)[1,1'-biphenyl]-4-yl]propyl]- (9CI) (CA INDEX NAME)

PAGE 1-A



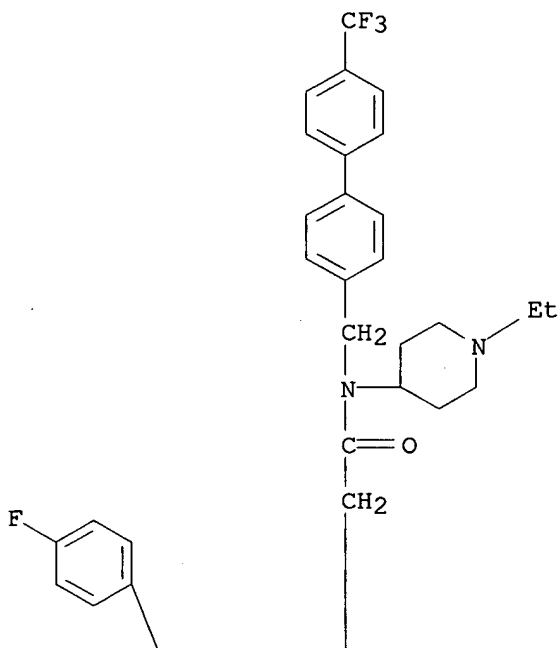
PAGE 2-A



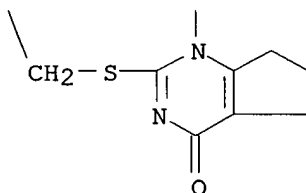
RN 356057-98-2 CAPLUS

CN 1H-Cyclopentapyrimidine-1-acetamide, N-(1-ethyl-4-piperidiny1)-2-[[(4-fluorophenyl)methyl]thio]-4,5,6,7-tetrahydro-4-oxo-N-[[4'-(trifluoromethyl)[1,1'-biphenyl]-4-yl]methyl]- (9CI) (CA INDEX NAME)

PAGE 1-A

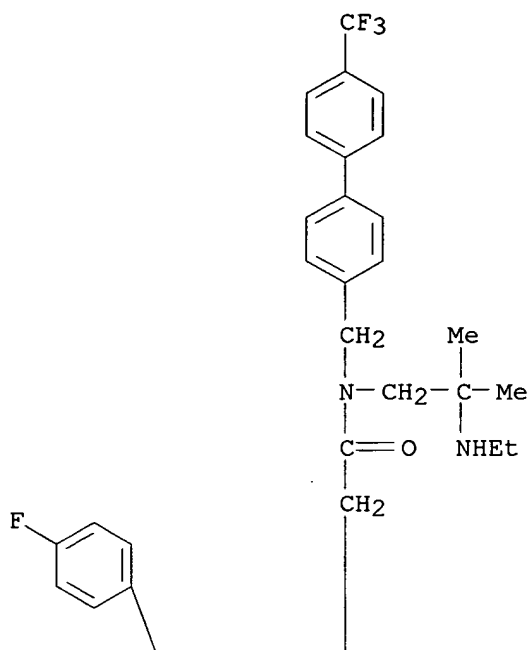


PAGE 2-A

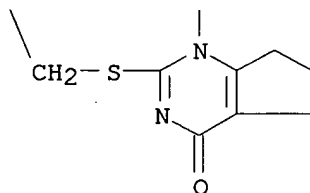


RN 356057-99-3 CAPLUS
 CN 1H-Cyclopentapyrimidine-1-acetamide, N-[2-(ethylamino)-2-methylpropyl]-2-[[(4-fluorophenyl)methyl]thio]-4,5,6,7-tetrahydro-4-oxo-N-[[4'-(trifluoromethyl)[1,1'-biphenyl]-4-yl]methyl]- (9CI) (CA INDEX NAME)

PAGE 1-A

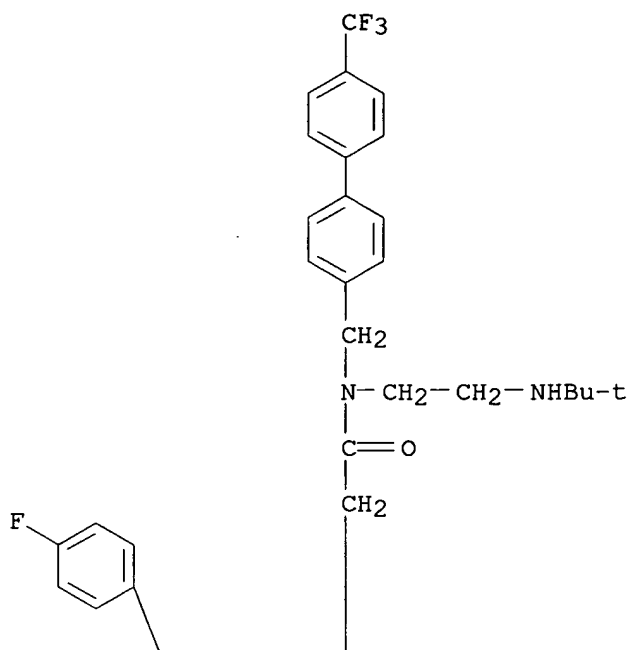


PAGE 2-A

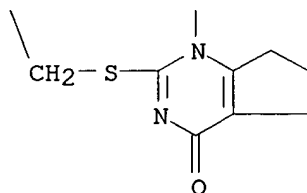


RN 356058-00-9 CAPLUS
 CN 1H-Cyclopentapyrimidine-1-acetamide, N-[2-[(1,1-dimethylethyl)amino]ethyl]-
 2-[[[(4-fluorophenyl)methyl]thio]-4,5,6,7-tetrahydro-4-oxo-N-[[4'-
 (trifluoromethyl)[1,1'-biphenyl]-4-yl]methyl]- (9CI) (CA INDEX NAME)

PAGE 1-A

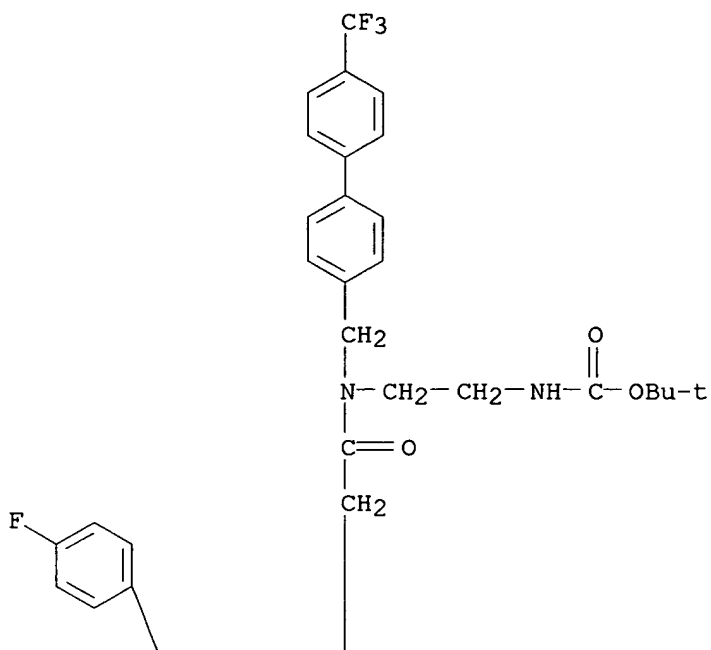


PAGE 2-A

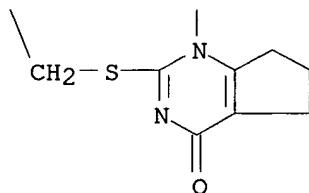


RN 356058-03-2 CAPLUS
 CN Carbamic acid, [2-[[[2-[[[4-(4-fluorophenyl)methyl]thio]-4,5,6,7-tetrahydro-4-oxo-1H-cyclopentapyrimidin-1-yl]acetyl][[4'-(trifluoromethyl)[1,1'-biphenyl]-4-yl]methyl]amino]ethyl]-, 1,1-dimethylethyl ester (9CI) (CA INDEX NAME)

PAGE 1-A

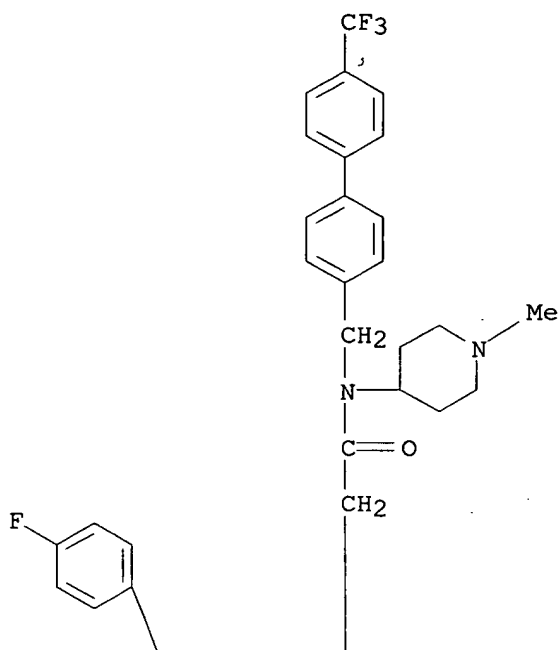


PAGE 2-A

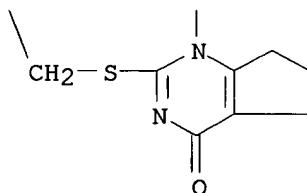


RN 356058-05-4 CAPLUS
 CN 1H-Cyclopentapyrimidine-1-acetamide, 2-[[[4-(4-fluorophenyl)methyl]thio]-
 4,5,6,7-tetrahydro-N-(1-methyl-4-piperidinyl)-4-oxo-N-[[4'-
 (trifluoromethyl)[1,1'-biphenyl]-4-yl]methyl]- (9CI) (CA INDEX NAME)

PAGE 1-A

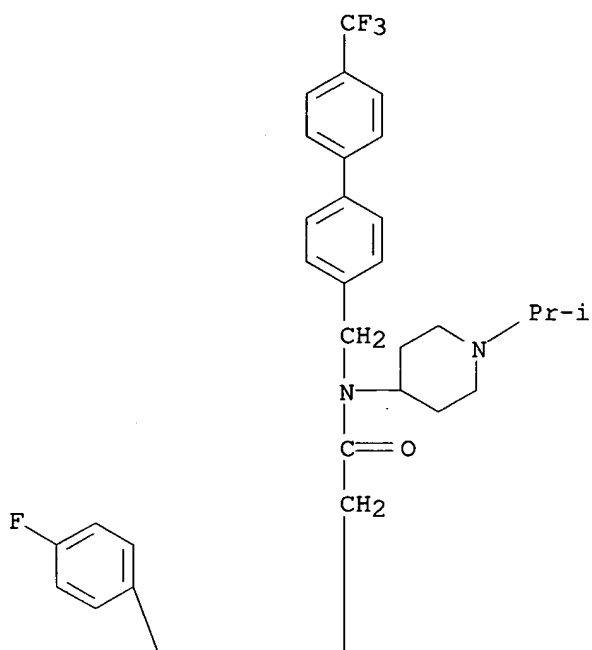


PAGE 2-A

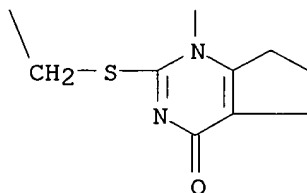


RN 356058-06-5 CAPLUS
 CN 1H-Cyclopentapyrimidine-1-acetamide, 2-[[[4-(4-fluorophenyl)methyl]thio]-
 4,5,6,7-tetrahydro-N-[1-(1-methylethyl)-4-piperidinyl]-4-oxo-N-[[4'-
 (trifluoromethyl)[1,1'-biphenyl]-4-yl]methyl]- (9CI) (CA INDEX NAME)

PAGE 1-A

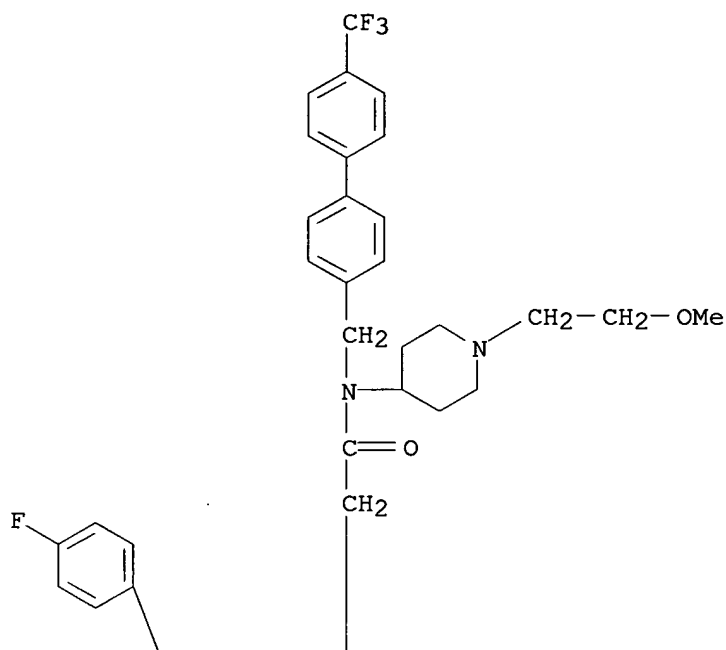


PAGE 2-A

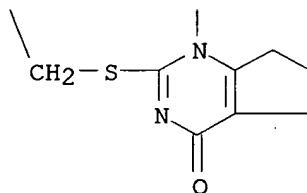


RN 356058-07-6 CAPLUS
 CN 1H-Cyclopentapyrimidine-1-acetamide, 2-[[[(4-fluorophenyl)methyl]thio]-
 4,5,6,7-tetrahydro-N-[1-(2-methoxyethyl)-4-piperidinyl]-4-oxo-N-[[4'-
 (trifluoromethyl)[1,1'-biphenyl]-4-yl]methyl]- (9CI) (CA INDEX NAME)

PAGE 1-A

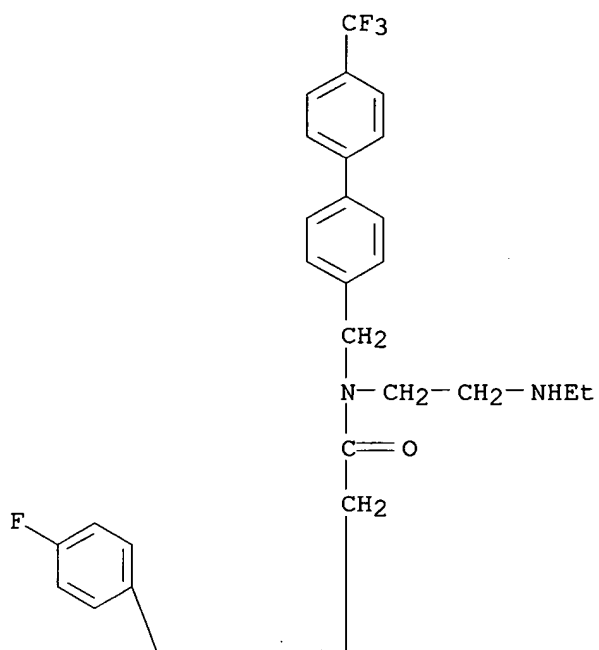


PAGE 2-A

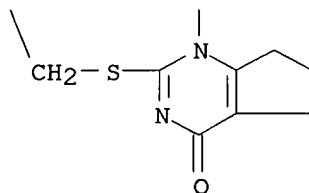


RN 356058-12-3 CAPLUS
 CN 1H-Cyclopentapyrimidine-1-acetamide, N-[2-(ethylamino)ethyl]-2-[[4-(4-fluorophenyl)methyl]thio]-4,5,6,7-tetrahydro-4-oxo-N-[[4'-(trifluoromethyl)[1,1'-biphenyl]-4-yl]methyl]- (9CI) (CA INDEX NAME)

PAGE 1-A



PAGE 2-A



RE.CNT 4 THERE ARE 4 CITED REFERENCES AVAILABLE FOR THIS RECORD
ALL CITATIONS AVAILABLE IN THE RE FORMAT

=> => d his

(FILE 'HOME' ENTERED AT 20:10:35 ON 07 DEC 2004)

FILE 'REGISTRY' ENTERED AT 20:10:39 ON 07 DEC 2004

L1 STRUCTURE UPLOADED

L2 1 S L1 SSS SAM

L3 29 S L1 SSS FUL

FILE 'CAPLUS' ENTERED AT 20:11:23 ON 07 DEC 2004

L4 6 S L3

FILE 'CAOLD' ENTERED AT 20:11:50 ON 07 DEC 2004

=> s 13

L5 0 L3

=> log y

COST IN U.S. DOLLARS

SINCE FILE

TOTAL

ENTRY

SESSION

FULL ESTIMATED COST

0.42

185.05

DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS)

SINCE FILE

TOTAL

ENTRY

SESSION

CA SUBSCRIBER PRICE

0.00

-4.20

STN INTERNATIONAL LOGOFF AT 20:12:01 ON 07 DEC 2004